FORM PTO-1449, Adapted

## LIST OF INFORMATION DISCLOSED BY APPLICANT

|  |   | RAD                                      | EMARK   | (Use seve  | eral sheets if nec           | essary)                           |          |                |                               |  |
|--|---|--|---|--|------------------------------|-----------------------------------|----------|----------------|-------------------------------|--|
|  |   |  | SERIAL NO.<br>09/491,50   |  | FILING DATE January 26, 2000 |                                   |          |                |                               |  |
| APPLICANT  Keith L. Black and Nagendra S. Ningaraj |   |  | arai  |  | GROUP<br>1632                |                                   |          |                |                               |  |
| Keitii L. E  | olack o   | and Magentin 3. Ming                     | ura)  | U.S. PA  | TENT DOCUM                   | IENTS                             |          |                | - 10                          |  |
| EXAMINER<br>INITIAL                                |   | DOCUMENT NUMBER                          | DATE  |  | NAME                         | NAME CLASS                        |          | SUBCLASS       | FILING DATE<br>IF APPROPRIATE |  |
| Ama  | AA  | 5,518,499                                | 18,499 05/21/06 Agar  |  | -                            |                                   |          |                |                               |  |
| Ama  | AB  | 5,767,160                                | 4   | 8 Kaesemeyer   |                              |                                   |          |                |                               |  |
|  |   |  |   |  | ding Author, Tit             |                                   |          |                |                               |  |
| Amo  | AC  |  |   | , Contribution of kca channel activation to hypoxic cerebrovasodilation does not involve NO, 4-48 (1998). ABSTRACT ONLY. |                              |                                   |          |                |                               |  |
| 1  | AD  | Barna, M., et                            | Barna, M., et al., Activation of type III nitric oxide synthase in astrocytes following a neurotropic viral infection, Virology, 223: 331-343 (1996).   |  |                              |                                   |          |                | rotropic viral                |  |
|  | AE  | effects in hum                           | an and rate   | platelets, ]   |                              | macol, 35(3):                     | 390-7 (2 | 000). ABST     | RACT ONLY.                    |  |
|  | AF  | meningitis, Br                           | effects in human and rate platelets, J Cardiovasc Pharmacol, 35(3):390-7 (2000). ABSTRACT ONLY. Boje, K.M., Inhibition of nitric oxide synthase attenuates blood-brain barrier disruption during experimental meningitis, Brain Research, 720:75-83 (1996). |  |                              |                                   |          |                |                               |  |
|  | AG  | microvasculat                            | Brandt, L., et al., Effects of topical application of calcium antagonist (nifedipine) on feline cortical pial microvasculature under normal conditions and in focal ischemia, Journal of Cerebral Blood Flow and Metabolism, 3:44-50 (1983).                |  |                              |                                   |          |                |                               |  |
|  | AH  |  | ., et al., Activators of soluble guarylate cyclase for treatment of male erectile dysfunction, al Journal of Impotence Research, 14:8-14 (2002).  |  |                              |                                   |          |                |                               |  |
|  | AI  | Bychkov, R., coronary arte               | R., et al. Calicum-activated potassium channels and nitrate-induced vasodilation in human arteries, J Pharacol Exp Therap, 285:293-8 (1998). ABSTRACT ONLY.   |  |                              |                                   |          |                |                               |  |
|  | AJ  |  | Chandran, S., et al., Nitric oxide: concepts, current perspectives and future therapeutic implications, Indian Journal of Pharmacology, 30:351-366 (1998).  |  |                              |                                   |          |                |                               |  |
|  | AK  |  | Chi, O.Z., et al. Effect of inhibition of nitric oxide synthase on blood-brain barrier transport in focal cerebral ischemia, Pharmacologylogy, 48:367-373 (1994).   |  |                              |                                   |          |                |                               |  |
|  | AL  |  | Cloughesy, T.F., et al., <i>Pharmacological blood-brain barrier modification for selective drug delivery</i> , Journal of Neuro-Oncology, 26:125-132 (1995).  |  |                              |                                   |          |                |                               |  |
|  | AM  |  | Feelisch, M., The use of nitric oxide donors in pharmacological studies, Naunyn-Schmiedeberg's Arch Pharmcol, 358:113-122 (1998).   |  |                              |                                   |          |                |                               |  |
|  | AN  | Fukao, M., et<br>mammalian co            | ao, M., et al., Cyclic GMP-dependent protein kindase activates cloned BKCa channels expressed in numalian cells by direct phophorylation at serine 1072, J Biol Chem, 274(16):10927-35 (1999).  |  |                              |                                   |          |                |                               |  |
|  | AO  |  | Fukumura, D., et al., Role of nitric oxide in angiogenesis and microcirculation in tumors, Cancer and Metastasis Reviews, 17:77-89 (1998).  |  |                              |                                   |          |                |                               |  |
| V  | AP  |  | le, P., et al., cGMP modulates basal and activated microvessel permeability independenty of [Ca2+]i, Am J Physiol, 274:H1865-74 (1998). ABSTRACT ONLY.  |  |                              |                                   |          |                |                               |  |
| Ama  | 1 AO Unam C.M. et al. Maintained vacadilatory response to gramakalim after inhibition of nitric oxide |  |   |  |                              |                                   |          | fnitric oxide  |                               |  |
| EXAMINE  | ER  | Inne-mari                                |   |  |                              | DATE CONSID                       | 2000     | 8/30/0         | 04                            |  |
| EXAMIN conforman                                   | ER: I   | Initial if reference considered. Include | lered, whethe<br>copy of this   | r or not cita<br>form with n   | tion is in conformation      | ance with MPEF<br>n to applicant. | 609; Dra | w line through | citation if not in            |  |
|  |   |  |   |  |                              |                                   |          |                |                               |  |

FORM PTO-1449, Adapted

## LIST OF INFORMATION DISCLOSED BY APPLICANT (Use several sheets if necessary)

| !   |   |                       |   |   |   |                                      |  |  |  |
|---|---|-----------------------|---|---|---|--------------------------------------|--|--|--|
| ATTY. DOCKET NO. SERIAL NO. 09/491,500                                    |   |                       |   |   | FILING DATE January 26, 2000  |                                      |  |  |  |
| APPLICANT Keith L. Black and Nagendra S. Ningaraj                         |   |                       |   | GROUI<br>1632   | •   |                                      |  |  |  |
|   |   | OTHER                 | REFERENCES (In  | cluding Autho   | uding Author, Title, Date, Pertinent Pages, Etc.)   |                                      |  |  |  |
|   |   |                       |   | e of cGMP in modulation of macromolecule permeability of aortic<br>2:H91-8 (1997). ABSTRACT ONLY. |   |                                      |  |  |  |
| AS Hongli, X., et al., Opening blood-b. malignant cerebral glioma, Chines |   |                       |   | <i>brain-barrier</i><br>ese Medical Jo  | rain-barrier by intracarotid infusion of papaverine in treatment of Medical Journal, 111(8):751-753 (1998). |                                      |  |  |  |
|   | AT Hurst, R.D., et al., Nitric oxide-induced perturbations in a cell culture mo Journal of Cellular Physiology, 167:89-94 (1996). |                       |   |   |   | odel of the blood-brain barrier,     |  |  |  |
|   | AU  |                       | Inamura, T., et al., Intracarotid histamine infusion increases blood tumour permeability in RG2 glioma, Neurological Research, 16:125-128 (1994).   |   |   |                                      |  |  |  |
|   | AV  |                       | Inamura, T., et al., Intracarotid infusion of RMP-7, a bradykinin analog: a method for selective drug delivery to brain tumors, J Neurosurg, 81:752-758 (1994).   |   |   |                                      |  |  |  |
|   | AW  |                       | Janigro, D., et al., Regulation of blood-brain barrier endothelial cells by nitric oxide, Circulation Research, 75:528-528 (1994).  |   |   |                                      |  |  |  |
|   | AX  |                       | Kimura, M., et al., Responses of human basilar and other isolated arteries to novel nitric oxide donors, <u>J.</u> Cardiovasc Pharmacol, 32: 695-701 (1998). ABSTRACT ONLY.   |   |   |                                      |  |  |  |
|   | AY  | Koesling, D., (1998). | Koesling, D., Modulators of soluble guanylyl cyclase, Naunyn-Schmiedeberg's Arch Pharmacol, 358:123-126   |   |   |                                      |  |  |  |
|   | AZ  |                       | Liu, Y., et al., Repeated, short-term ischemia augments bradykinin-mediated opening of the blood-tumor barrier in rats with RG2 glioma, Neurological Research, 23:631-639 (2001).   |   |   |                                      |  |  |  |
|   | BA  | Lohse, M.J.,          | Lohse, M.J., et al., <i>Pharmacology of NO:cGMP signal transduction</i> , Naunyn-Schmiedeberg's Arch Pharmcol, 358:111-112 (1998).  |   |   |                                      |  |  |  |
|   | ВВ  |                       | Matukado, T., et al., Selective Increase in Blood Tumor Permeability by Calcium Antagonists in Transplanted Brain Tumors, Acta Neurochir, 60: 403-405 (1994).   |   |   |                                      |  |  |  |
|   | BC  |                       | Mayer, B., et al., Nitric oxide synthases: catalytic function and progress toward selective inhibition, Naunyn-Schmiedeberg's Arch Pharmcol, 358:127-133 (1998).  |   |   |                                      |  |  |  |
|   | BD  |                       | Mayhan, W.G., Role of nitric oxide in histamine-induced increases in permeability of the blood-brain barrier, Brain Research, 743:70-76 (1996).   |   |   |                                      |  |  |  |
|   | BE  | Mayhan, W.(           | Mayhan, W.G., et al., Glutamate-induced disruption of the blood-brain barrier in rats, Stroke, 27:965-970   |   |   |                                      |  |  |  |
| V   | BF  |                       | Nakano, S., et al., Increased brain microvessel permeability after intracarotid bradykinin infusion is mediated by nitric oxide, Cancer Research, 56:4027-4031 (1996).  |   |   |                                      |  |  |  |
| Ama   | BG  | of Neurologic         | Ningaraj, N.S., et al., Role of ATP-sensitive K+ channels in blood-brain tumor barrier permeability, Congress of Neurological Surgeons Annual Meeting, 50 <sup>th</sup> Anniversary Celebration, September 23-28, 2000, Henry B. Gonzalez Convention Center, San Antonio, Texas, ABSTRACT No. 4309, p. 215. |   |   |                                      |  |  |  |
| EXAMINER Anne-Marie Falk  |   |                       |   |   | DATE CONSIDERED   |                                      |  |  |  |
|   |   |                       | dered, whether or not o   |   |   | Draw line through citation if not in |  |  |  |

FORM PTO-1449, Adapted

## Wise several sheets if necessary)

| ATTY. DOCKET NO. SERIAL NO. 09/491,500            |  |   |   | FILING DATE January 26, 2000   |  |  |  |  |
|---|--|---|---|--|--|--|--|--|
| APPLICANT Keith L. Black and Nagendra S. Ningaraj |  |   | garaj   | GROUP<br>1632  |  |  |  |  |
|   |  | OTHER   | REFERENCES (In  | ncluding Author, Title, Date, Pertinent Pages, Etc.)   |  |  |  |  |
| permeability, Congress of Neurologic              |  |   |   | ent K+ channels are a key regulatory of blood-brain tumor barrier cal Surgeons Annual Meeting, 50 <sup>th</sup> Anniversary Celebration, September ovention Center, San Antonio, Texas, ABSTRACT No.428, p. 219. |  |  |  |  |
|   | BI Ningaraj, N.S., et al., Nitric oxide donors increase blood-brain tumor barrier permeability via Kca chan Society for Neuroscience, 30th Annual Meeting, New Orleans, LA, November 4-9, 2000, 26 Part 1, p. 33 ABSTRACT No. 126.8. |   |   |  |  |  |  |  |
|   | BJ   |   | Ningaraj, N.S., et al., Regulation of blood-brain tumor barrier permeability by calcium-activated potassium channels, The Journal of Pharmacology, June 2002, 301: 838-851                                |  |  |  |  |  |
|   | BK   | Pardrige, W. (1992).  | Pardrige, W., et al., Blood -brain barrier and new approaches to drug delivery, West J Med, 156:281-286 (1992).   |  |  |  |  |  |
|   | BL   |   | Robertson, B.E., et al., cGMP-dependent protein kinase activates Ca-activated K channels in cerebral artery smooth muscle cells, Am J Physiol, 265:C299-C303 (1993).                                      |  |  |  |  |  |
|   | BM   |   | Sobey, C.G., et al., Inhibitory effect of 4-aminopyridine on responses of the basilar artery to nitric oxide, Br J Pharmacol, 126:1437-43 (1999). ABSTRACT ONLY.  |  |  |  |  |  |
|   | BN   |   | Salom, J.B., et al., Relaxant effects of sodium nitroprusside and NONOates in rabbit basilar artery,<br>Pharmacology, 57:79-97 (1998). ABSTRACT ONLY.   |  |  |  |  |  |
|   | ВО   |   | Salom, J.B., et al., Comparative relaxant effects of the NO donors sodium nitroprusside, DEA/NO and SPER/NO in rabbit carotid arteries, Gen Pharmacol, 32:75-59 (1999). ABSTRACT ONLY.                    |  |  |  |  |  |
|   | BP   |   | Salom, J.B., et al., Relexant effects of sodium nitroprusside and NONates in goat middle cerebral artery: delayed impairment of global ischemia-reperfusion, Nitric Oxide, 3:85-93 (1999). ABSTRACT ONLY. |  |  |  |  |  |
|   | BQ   |   | Shukla, A., et al., Nitric oxide-dependent blood-brain barrier permeability alteration in the rat brain,<br>Experientia, 52:136-140 (1996).   |  |  |  |  |  |
|   | BR   |   | Smolenski, A., et al., Functional analysis of cGMP-dependent protein kinases I and II as mediators of NO/cGMP effects, Naunyn-Schmiedeberg's Arch Pharmacol, 358:134-138.                                 |  |  |  |  |  |
|   | BS   |   | Sugita, M., et al., Cyclic GMP-specific phosphodiesterase inhibition and intracarotid bradykinin infusion enhances permeability in brain tumors, Cancer Research, 58:914-920 (1998).                      |  |  |  |  |  |
| V   | BT   |   | Takayasu, M., et al., Effects of calcium antagonists on intracerebral penetrating arteriolesi in rats, <u>I</u> Neurosurg, 69:104-109 (1988).   |  |  |  |  |  |
| Amo   | BU   | Uchida, M., et al., Overexpression of bradykinin type 2 receptors on glioma cells enhances bradykinin-<br>mediated blood-brain tumor varrier permeability increase, Neurological Research, 24:739-745 |   |  |  |  |  |  |
| EXAMINER Anne-Marie Falk DATE CONSIDERED 8/30/04  |  |   |   |  |  |  |  |  |
| EXAMIN  | ER: In   | itial if reference consi  | dered, whether or not c   | citation is in conformance with MPEP 609; Draw line through citation if not in   |  |  |  |  |

conformance and not considered. Include copy of this form with next communication to applicant.

## CUse several sheets if necessary)

| ATTY. DOCKET  | ΓNO.   | SERIAL NO.<br>09/491,500   |               | FILING DATE January 26, 2000                          |  |  |  |  |
|---|--|--|---------------|---|--|--|--|--|
| APPLICANT Keith L. Black and Nagendra S. Ningaraj   |  |  | GROUP<br>1632 |   |  |  |  |  |
|   | ОТНЕ   | R REFERENCES (In   | cluding Autho | r, Title, Date, Pertinent Pages, Etc.)                |  |  |  |  |
| Uchida, M., et al., Cyclic GMP-dependent blood-brain tumor ba<br>GMP-dependent protein kinase, Congress of Neurological Surge<br>Celebration, September 23-28, 2000, Henry B. Gonzalez Conver<br>No. 440, p. 220. |  |  |               | eurological Surgeons Annual Meeting, 50th Anniversary |  |  |  |  |
| \ B   |  | Vodovotz, Y., et al., Regulation of transforming growth factor beta 1 by nitric oxide, Cancer Res, 59:2142-9 (1999). ABSTRACT ONLY.                                    |               |   |  |  |  |  |
| B   | BX Yukabu, M.A., Hematoma-induced enhanced cerebral vasoconstriction to leukotriene C4 and endothe piglets: role of prostanoids, Pediatr Res, 38:119-23 (1995). ABSTRACT ONLY. |  |               |   |  |  |  |  |
| √ B   | Y Tocris We 5/31/00.   | Tocris Web Page, <a href="http://www.tocris.com/cat/nodonorstxt.html">http://www.tocris.com/cat/nodonorstxt.html</a> No Donors/Precursors, pp.1-2, Downloaded 5/31/00. |               |   |  |  |  |  |
| Amalb   | Z   Sigma-Ald  | Sigma-Aldrich Web page, http://vsearch.sial.com/search 97cgi/s97-cgi, p.1, downloaded 5/31/00.   |               |   |  |  |  |  |
| EXAMINER Ame-marie Talk   |  |  |               | DATE CONSIDERED 8/30/04                               |  |  |  |  |
|   |  |  |               |   |  |  |  |  |

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.